

THE CLAIMS DEFINING THE INVENTION ARE AS FOLLOWS:

1. A method of diagnosing and/or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including:
 - a) determination of IgG2 anti-*H. pylori* antibody level in the subject;
 - 5 b) comparison of the IgG2 anti-*H. pylori* antibody level with a predetermined control IgG2 anti-*H. pylori* antibody level, wherein a reduction in the level of IgG2 anti-*H. pylori* antibody in the subject compared to the control indicates the presence and/or increased risk of developing gastric cancer.
2. A method of diagnosing and/or determining the risk of developing gastric cancer
10 in a subject with a *Helicobacter* infection, including:
 - a) determination of γ IFN level in the subject;
 - b) comparison of the γ IFN level with a predetermined control γ IFN level, wherein a reduction in the level of γ IFN in the subject compared to the control indicates the presence and/or increased risk of developing gastric cancer.
- 15 3. A method of diagnosing and/or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including:
 - a) determination of IL-4 level in the subject;
 - b) comparison of the IL-4 level with a predetermined control IL-4 level, wherein
an elevation in the level of IL-4 in the subject compared to the control indicates the
20 presence and/or increased risk of developing gastric cancer.
4. A method of diagnosing and/or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including a combination of a method

- 15 -

according to claim 1 and/or a method according to claim 2 and/or a method according to claim 3.

- 5 5. A method of diagnosing and/or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including a combination of a method according to claim 2 and a method according to claim 3.
6. A method according to any one of claims 1 to 6 wherein the *Helicobacter* infection is a *Helicobacter pylori* infection.
7. A method according to any one of claims 1 to 7 wherein the IgG2 anti-*H. pylori* antibody, γ IFN and/or IL-4 levels are determined by detection of the levels in a sample
10 of biological fluid.
8. A method according to claim 7 wherein the biological fluid is blood.
9. A method according to claim 7 wherein the biological fluid is saliva.
10. A method according to claim 7 wherein the biological fluid is gastric fluid.
11. A method according to any one of claims 1 to 10 wherein the measurement of
15 IgG2 anti-*H. pylori* antibody, γ INF and/or IL-4 either simultaneously provides, or can be performed simultaneously with, a method which provides an indication of *H. pylori* status.
12. A method according to any one of claims 1 to 11 wherein the IgG2 anti-*H. pylori* antibody, γ IFN and/or IL-4 are detected by a near-subject assay.
- 20 13. A method according to any one of claims 1 to 11 wherein the assay is a laboratory-based test.
14. A method according to claim 12 or claim 13 wherein the assay is an antibody assay.

- 16 -

15. A method according to claim 14 wherein the antibody assay is an ELISA.
16. A method of predicting the risk of, and/or diagnosing, gastric cancer in a subject having a *Helicobacter* infection by
- a) determining the frequency of IgG2 anti-*H.pylori* antibody- and/or γ IFN- and/or
- 5 IL-4-producing cells in the subject's blood; and
- b) comparison of the frequency of IgG2 anti-*H.pylori* antibody- and/or γ IFN- and/or IL-4-producing cells in the subject's blood with a predetermined control level, wherein a reduction in the level of IgG2 anti-*H.pylori* antibody- and/or γ IFN-producing cells and/or an elevation in IL-4-producing cells in the subject's blood indicates the
- 10 presence and/or increased risk of developing gastric cancer.
17. A method according to claim 16 wherein the blood is purified to provide an enriched white blood cell population.
18. A method according to claim 17 wherein the white blood cell population is further fractionated to obtain specific cell populations.
- 15 19. A method according to any one of claims 16 to 18 wherein the IgG2 anti-*H.pylori* antibody- and/or γ IFN- and/or IL-4-producing cells are stimulated with *H. pylori* antigen prior to measurement of IgG2 anti-*H.pylori* antibody and/or γ IFN and/or IL-4.
20. A method of predicting the risk of, and/or diagnosing, gastric cancer in a subject having a *Helicobacter* infection by
- 20 a) determining the frequency of IgG2 anti-*H.pylori* antibody and/or γ IFN and/or IL-4-producing cells in the subject's gastric mucosa; and
- b) comparison of the frequency of IgG2 anti-*H.pylori* antibody and/or γ IFN and/or IL-4-producing cells in the subject's gastric mucosa with a predetermined control level,

- 17 -

wherein a reduction in the level of IgG2 anti-*H.pylori* antibody- and/or γ IFN-producing cells and/or an elevation in IL-4-producing cells in the subject's gastric mucosa indicates the presence and/or increased risk of developing gastric cancer.

21. A method according to claim 20 wherein the cells are derived from a biopsy
5 sample.

22. A method according to claim 20 or claim 21 wherein of IgG2 anti-*H.pylori* antibody and/or γ IFN and/or IL-4-producing cells are detected by flow cytometry.